









computer program with a GAP creation penalty of 5.0 and GAP extension penalty of 0.3, and is capable of initiating replication; and (2) replication initiating sequence which hybridises under low stringency conditions with (i) the nucleic acid sequence of SEQ ID NO:1 or SEQ ID NO:2, or (ii) the respective complementary strands, wherein the low stringency conditions are defined by prehybridization and hybridization at 42°C in 5x SSPE, 0.3% SDS, 200 mg/ml sheared and denatured salmon sperm DNA, and 25% formamide, and wash conditions are defined at 50°C for 30 minutes in 2x SSC, 0.2% SDS; and

- (ii) a polynucleotide sequence of interest having or encoding a desired activity or function characteristic, wherein there are vectors in the population that vary from other vectors in the population by carrying different versions of the polynucleotide sequence of interest;
- (b) cultivating the cells in the presence of an effective amount of a selective agent or the absence of an appropriate selective agent.

32. (Currently amended) The method of claim 31, further comprising the steps of:

- (c) selecting or screening for one or more transformants expressing the desired activity or function characteristic; and
- (d) isolating the transformant(s) of interest.